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HARPOLD, *et al.*  
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IN THE CLAIMS:

Please cancel claim 57, 58, 61, 63, 67, 71, 73, 74, 76-78, 86-90 and 99 without prejudice or disclaimer.

Please add claims 100-112, which replace and amend cancelled claims as follows:

— 100. An isolated nucleic acid, comprising the sequence of nucleotides set forth in SEQ ID No. 1.—

— 101. An isolated nucleic acid, comprising the sequence of nucleotides set forth in SEQ ID No. 3.—

— 102. An isolated nucleic acid, comprising the sequence of nucleotides set forth in SEQ ID No. 5.—

— 103. An isolated nucleic acid, comprising the sequence of nucleotides set forth in SEQ ID No. 7.—

— 104. An isolated nucleic acid molecule encoding a beta2 subunit of a human neuronal nicotinic acetylcholine receptor, comprising the sequence of nucleotides set forth in SEQ ID No. 9.—

— 105. A plasmid having all of the identifying characteristics of the plasmid deposited under ATCC Accession No. 68278.—

— 106. A plasmid having all of the identifying characteristics of the plasmid deposited under ATCC Accession No. 68279.—

— 107. The cell of claim 59 that is a bacterial or eukaryotic cell.—

— 108. The eukaryotic cell of claim 107 that is a mammalian cell, yeast cell or amphibian oocyte.—

— 109. An isolated cell, comprising the nucleic acid of claim 60.—

— 110. The cell of claim 109 that is a bacterial or eukaryotic cell.—

— 111. The eukaryotic cell of claim 110 that is a mammalian cell, yeast cell or amphibian oocyte.—

— 112. The eukaryotic cell of claim 111 that expresses a nicotinic acetylcholine receptor comprising a subunit encoded by the nucleic acid.—

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Please amend claims 53, 55, 56, 59, 60, 62, 63, 66, 68, 70, 72, 82-84, 91, 92 and 98 as follows:

53. (Amended) An isolated nucleic acid molecule, comprising a sequence of nucleotides encoding [an alpha2 subunit of a human nicotinic acetylcholine receptor that is encoded by nucleic acid comprising the sequence of nucleotides set forth SEQ ID No. 1 and 3, an alpha3 subunit of a human nicotinic acetylcholine receptor that is encoded by nucleic acid comprising the sequence of nucleotides set forth in SEQ ID No. 5 and 7 or] a beta2 subunit of a human neuronal nicotinic acetylcholine receptor[ that is encoded by nucleic acid comprising the sequence of nucleotides set forth in SEQ ID No. 9].

55. (Amended) [A] An isolated and purified [substantially pure subunit of the] human neuronal nicotinic acetylcholine receptor subunit encoded by the alpha3-encoding nucleic acid in a plasmid having all of the identifying characteristics of HnAChRa3 deposited under ATCC Accession No. 68278.

56. (Amended) [A] An isolated and purified [substantially pure subunit of the] human neuronal nicotinic acetylcholine receptor subunit encoded by the beta2-encoding nucleic acid [of claim 53]in a plasmid having all of the identifying characteristics of HnAChRa3 deposited under ATCC Accession No. 68279.

59. (Amended) An isolated cell, comprising [containing any one or more of] the nucleic [acids of claim 53] acid molecule of claim 80.

60. (Amended) An isolated cell, comprising [containing one or more of] the nucleic [acids of claim 54] acid molecule of claim 81.

62. (Amended) The cell of claim [59] 60 that is a eukaryotic cell.

63. The cell of claim 59 that is a bacterial cell, mammalian cell, yeast cell or amphibian oöcyte.

66. The cell of claim 59 further comprising a nucleic acid molecule that encodes a beta subunit of a human nicotinic acetylcholine receptor, wherein the beta subunit comprises a sequence of amino acids encoded by SEQ ID No. 9.

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68. (Twice amended) A method for screening compounds for activity as nicotinic acetylcholine receptor agonists or antagonists, said method comprising:

contacting [cells] a cell of claim 59 with a test compound, and thereafter monitoring nicotinic acetylcholine receptor activity of the cells by monitoring the performance of the [cells] cell by measuring a performance parameter selected from the group consisting of the flux of ions through the [membranes] membrane of the [cells] cell, nicotine binding to nicotinic acetylcholine receptors of the cell, or the electrophysiological response of the cells, wherein[:

the cells contain one or more nucleic acids comprising a sequence of nucleotides (i) encoding an alpha2 subunit of a human nicotinic acetylcholine receptor and comprising a sequence of nucleotides set forth SEQ ID No. 1 and 3, (ii) encoding an alpha3 subunit of a human nicotinic acetylcholine receptor [and comprising a sequence of nucleotides set forth in SEQ ID No. 5 and 7, or (iii) encoding a beta2 subunit of a human neuronal nicotinic acetylcholine receptor and comprising a sequence of nucleotides set forth in SEQ ID No. 9; and] the [cells express] cell expresses a nicotinic acetylcholine receptor that contains a [one or more subunits] subunit encoded by the nucleic [acids] acid molecule.

70. (Amended) The method of claim 68, wherein the [alpha subunit is an alpha2 subunit encoded by a sequence of nucleotides having the restriction map of the DNA encoding the human alpha2 subunit set forth in Figure 1 or an] alpha3 subunit encoded by a sequence of nucleotides having the restriction map of the DNA encoding the human alpha3 subunit set forth in Figure 2, and the beta subunit is encoded by a sequence of nucleotides having the restriction map of the DNA encoding the human beta2 subunit set forth in Figure 3] the cell further comprises DNA encoding a beta2 subunit of a human nicotinic

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acetylcholine receptor comprising a sequence of amino acids encoded by SEQ ID No. 9.

72. (Amended) A method of making cells having neuronal nicotinic acetylcholine receptor activity, comprising:

(a) introducing one or more nucleic acid molecules that encode(s) at least one alpha subunit of a neuronal nicotinic acetylcholine receptor and at least one beta subunit of a neuronal nicotinic acetylcholine receptor, eukaryotic cells, wherein the nucleic acid encoding an [α] alpha subunit comprises [the] a sequence of amino acids encoded by [SEQ ID No. 1 or 3, or comprises the sequence of amino acids encoded by SEQ ID No. 5 or 7,] the alpha3-encoding nucleic acid that is isolated from a plasmid having all of the identifying characteristics of HnAChRa3 deposited under ATCC Accession No. 68278, and the nucleic acid encoding the beta subunit comprises [the] a sequence of amino acids encoded by SEQ ID No. 9;

(b) selecting cells from (a) that express the alpha or the beta encoding nucleic acid or express the alpha and beta subunit-encoding nucleic acid; and

(c) detecting neuronal nicotinic acetylcholine receptor activity in the selected cells, wherein the activity is mediated by a receptor containing one or more of the alpha and beta subunits encoded by said introduced nucleic acid molecules.

79. (Amended) An isolated [Isolated] nucleic acid[ of claim 57] molecule, comprising the alpha2-encoding nucleic acid open reading frame that is isolated from a plasmid having all of the identifying characteristics of HnAChRa2 deposited under ATCC Accession No. 68277.

80. (Amended) [Isolated nucleic acid of claim 57, comprising] An isolated nucleic acid molecule, comprising a sequence of nucleotides encoding an alpha3 subunit of a human nicotinic acetylcholine receptor that is encoded by the alpha3-encoding nucleic acid that is isolated from a plasmid having all of the

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identifying characteristics of HnAChR $\alpha$ 3 deposited under ATCC Accession No. 68278.

81. (Amended) [Isolated] An isolated nucleic acid molecule, [of claim 57] comprising a sequence of nucleotides encoding a beta2 subunit of a human nicotinic acetylcholine receptor that is encoded by the beta2-encoding nucleic acid that is isolated from a plasmid having all of the identifying characteristics of HnAChR $\beta$ 2 deposited under ATCC Accession No. 68279 or the open reading frame set forth in SEQ ID No. 9.

84. (Twice amended) An isolated and purified subunit of a human nicotinic acetylcholine receptor encoded by the beta2-encoding DNA set forth in SEQ ID NO. 9 [nucleic acid of claim 81].

82. (Amended) An isolated and purified subunit of a human nicotinic acetylcholine receptor encoded by the nucleic acid of claim 79.

83. (Amended) An isolated and purified subunit of a human nicotinic acetylcholine receptor encoded by the nucleic acid of claim 80.

84. (Amended) An isolated and purified subunit of a human nicotinic acetylcholine receptor encoded by the nucleic acid of claim 81.

91. (Amended) The cell of claim [61] 59, further comprising [that additionally contains] a reporter gene expression construct; and the reporter gene expression construct comprises:  
a transcriptional control element, and  
a reporter gene encoding a transcriptional and/or translational product;  
the transcriptional control element, in said cell, is responsive to an intracellular condition that occurs when a human neuronal nicotinic acetylcholine receptor interacts with a compound having agonist or antagonist activity with respect to said receptor;

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said product can be, directly or indirectly, detected; and  
the reporter gene is in operative association with said transcriptional  
control element.

92. (Amended) A method for screening test compounds for activity as  
nicotinic acetylcholine receptor agonists or antagonists, comprising:

comparing the difference in the amount of transcription of a reporter gene  
in the cells of claim 91 in the presence of the compound with the amount of  
transcription in the absence of the compound or with the amount of  
transcription in the control cells that do not express nicotinic acetylcholine  
receptors, but contain the reporter gene expression construct, wherein  
compounds that exhibit activity as agonists or antagonists are identified[,  
wherein:

the cells contain one or more nucleic acids comprising a sequence of  
nucleotides (i) encoding an alpha2 subunit of a human nicotinic acetylcholine  
receptor and comprising a sequence of nucleotides set forth SEQ ID No. 1 and  
3, (ii) encoding an alpha3 subunit of a human nicotinic acetylcholine receptor  
and comprising a sequence of nucleotides set forth in SEQ ID No. 5 and 7, or  
(iii) encoding a beta2 subunit of a human neuronal nicotinic acetylcholine  
receptor and comprising a sequence of nucleotides set forth in SEQ ID No. 9;

the cells also contain a reporter gene expression construct; the  
reporter gene expression construct comprises:

a transcriptional control element, and

a reporter gene encoding a transcriptional and/or translational product;

the transcriptional control element, in said cell, is responsive to an  
intracellular condition that occurs when a human neuronal nicotinic acetylcholine  
receptor interacts with a compound having agonist or antagonist activity with  
respect to said receptor;

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said product can be, directly or indirectly, detected; and  
the reporter gene is in operative association with said transcriptional  
control element; and

the cells express a nicotinic acetylcholine receptor that contains one or  
more subunits encoded by the nucleic acids].

96. An isolated nucleic acid molecule, comprising the sequence of  
nucleotides set forth in SEQ ID No. 9.

97. (Amended) An isolated [Isolated] nucleic acid molecule, comprising  
a sequence of nucleotides that encodes a beta2 subunit of a human nicotinic  
acetylcholine receptor, wherein the beta2 subunit[ that] comprises [a] the  
sequence of amino acids encoded by the sequence of nucleotides set forth as  
nucleotides 1-1521 in SEQ ID No. 9.

98. (Amended) An isolated and purified beta2 subunit of a human  
nicotinic acetylcholine receptor encoded by [the] a nucleic acid molecule, [of  
claim 97] comprising a sequence of nucleotides that encodes a beta2 subunit of  
a human nicotinic acetylcholine receptor, wherein the beta2 subunit comprises a  
sequence of amino acids encoded by the sequence of nucleotides set forth in  
SEQ ID No. 9.

#### REMARKS

A Notice of Appeal and a check for the part of the requisite fees for an extension of time and the Notice of Appeal accompany this Amendment. Any deficiency in these fees, including those for the extension of time, the Notice of Appeal and additional independent claims, that may be due may be charged to Deposit Account No. 02-4070. If a Petition for extension of time is needed, this paper is to be considered such Petition.

Claims 53, 55, 56, 59, 60, 62, 66, 68, 70, 72, 79-85, 91-98 and 100-112 are presently pending in this application (assuming entry of the instant Amendment). Claims 79, 81, 85 are allowable and claim 99, which has been replaced by claims 101-104 was orally indicated to be allowable (the Office